

Application No.: 09/701,586
Inventor: Kock et al.
Reply to Office Action of May 22, 2008
Docket No.: 49100

Amendments to the Claims:

The following Listing of Claims will replace all prior versions, and listings, of the claims in the above-identified application.

Listing of Claims

1. (currently amended) An isolated and purified-poly(ADP-ribose) polymerase (PARP) homolog comprising human PARP2 (SEQ ID NO: 2) or a functional equivalent thereof which is at least 95% 85% homologous to human PARP2 (SEQ ID NO: 2) thereto, exhibits poly(ADP-ribose)-synthesizing activity, and has an amino acid sequence which
 - a) has a functional NAD⁺ binding domain comprising the sequence motif $PX_n(S/T)GX_3GKGIYFA$ (SEQ ID NO:11) in which n is an integral value from 1 to 5, and the X radicals are, independently of one another, any amino acid;
and
b) lacks a zinc finger sequence motif of the formula $CX_2CX_mHX_2C$ (SEQ ID NO:30) in which m is an integral value of 28 or 30, and the X radicals are, independently of one another, any amino acid.
2. (currently amended) The PARP homolog as claimed in claim 1, wherein the functional NAD⁺ binding domain comprises the following sequence motif:
 $(S/T)XGLR(I/V)XPX_n(S/T)GX_3GKGIYFA$ (SEQ ID NO:12)
in which n is an integral value from 1 to 5, and the X radicals are, independently of one another, any amino acid.
3. (currently amended) The PARP homolog as claimed in claim 1, further comprising the [[part-]] sequence motif:

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LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15)

in which the X radicals are, independently of one another, any amino acid.

4-32. (canceled)

33. (currently amended) The PARP homolog as claimed in claim 1, wherein the functional NAD⁺ binding domain comprises the following sequence ~~motif~~:

LLWHG(S/T)X₇IL(S/T)XGLR(I/V)XPX_n(S/T)GX₃GKGIYFAX₃SKSAXY (SEQ ID NO:13)

in which n is an integral value from 1 to 5, and
the X radicals are, independently of one another, any amino acid.

34. (currently amended) The PARP homolog as claimed in claim 1, further comprising [[part-]] sequence: ~~motif~~

AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16)

in which the X radicals are, independently of one another, any amino acid.

35. (currently amended) The PARP homolog as claimed in claim 1, further comprising [[part-]] sequence: ~~motif~~

XL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17)

in which the X radicals are, independently of one another, any amino acid.

36. (currently amended) The PARP homolog as claimed in claim 1, further comprising [[part-]] sequence: ~~motif~~

FYTXIPHFGX₃PP (SEQ ID NO:18)

in which the X radicals are, independently of one another, any amino acid.

37. (currently amended) The PARP homolog as claimed in claim 1, further comprising

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[[part-]] sequence: ~~motif~~

$\text{KX}_3\text{LX}_2\text{LXDIEXAX}_2\text{L}$ (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid.

38. (currently amended) An isolated poly(ADP-ribose) polymerase (PARP) homolog comprising human PARP2 (SEQ ID NO: 2) or a functional equivalent thereof which is at least 95% 85% homologous to human PARP2 (SEQ ID NO: 2) thereto, exhibits poly(ADP-ribose)-synthesizing activity, and has an amino acid sequence which

- a) has a functional NAD^+ binding domain comprising the sequence ~~motif~~
 $\text{PX}_n(\text{S/T})\text{GX}_3\text{GKGIYFA}$ (SEQ ID NO:11)
in which n is an integral value from 1 to 5, and the X radicals are, independently of one another, any amino acid;

and

- b) lacks a zinc finger sequence ~~motif~~ of the formula
 $\text{CX}_2\text{CX}_m\text{HX}_2\text{C}$ (SEQ ID NO:30)
in which m is an integral value of 28 or 30, and the X radicals are, independently of one another, any amino acid
further comprising a leucine zipper-like sequence ~~motif~~:
 $(\text{L/V})\text{X}_6\text{LX}_6\text{LX}_6\text{L}$ (SEQ ID NO: 14)
wherein X radicals are, independently of one another, any amino acid.

39. (currently amended) The PARP homolog as claimed in claim 38, further comprising at least one of the following [[part-]] sequences ~~motifs~~:

$\text{LX}_9\text{NX}_2\text{YX}_2\text{QLLX(D/E)X}_{10/11}\text{WGRVG}$ (SEQ ID NO: 15),
 $\text{AX}_3\text{FXKX}_4\text{KTXNXWX}_5\text{FX}_3\text{PXK}$ (SEQ ID NO:16),
 $\text{QXL(I/L)X}_2\text{IX}_9\text{MX}_{10}\text{PLGKLX}_3\text{QIX}_6\text{L}$ (SEQ ID NO:17),
 $\text{FYTXIPHFGX}_3\text{PP}$ (SEQ ID NO:18), and
 $\text{KX}_3\text{LX}_2\text{LXDIEXAX}_2\text{L}$ (SEQ ID NO:19)

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in which the X radicals are, independently of one another, any amino acid.

40. (currently amended) The PARP homolog as claimed in claim 38, further comprising
[[part-]] sequences ~~motifs~~:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15)
AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),
QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),
FYTXIPHXFGX₃PP (SEQ ID NO:18), and
KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid.

41. (currently amended) The PARP homolog as claimed in claim 38, further comprising
[[part-]] sequences ~~motifs~~:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15)
AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),
QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),
FYTXIPHXFGX₃PP (SEQ ID NO:18), and
KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid, wherein

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO:15)

is closest to the N terminus.

42. (currently amended) The PARP homolog as claimed in claim 1, further comprising
[[part-]] sequences ~~motifs~~:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15)
AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),
QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),
FYTXIPHXFGX₃PP (SEQ ID NO:18), and

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KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid.

43. (currently amended) The PARP homolog as claimed in claim 1₂, further comprising [[part-]] sequences ~~motifs~~:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15)

AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),

QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),

FYTXIPHXFGX₃PP (SEQ ID NO:18), and

KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid, wherein

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO:15)

is closest to the N terminus.

44. (currently amended) The PARP homolog as claimed in claim 1₂, further comprising at least one of the following:

GX₃LXEVALG (SEQ ID NO: 20),

GX₂SX₄GX₃PX_aLXGX₂V (SEQ ID NO: 21), and

E(Y/F)X₂YXYX₃QXYLL (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid.

45. (currently amended) The PARP homolog as claimed in claim 1₂, further comprising

GX₃LXEVALG (SEQ ID NO: 20),

GX₂SX₄GX₃PX_aLXGX₂V (SEQ ID NO: 21), and

E(Y/F)X₂YX₃QX₄YLL (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid.

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46. (currently amended) The PARP homolog as claimed in claim 1, further comprising
GX₃LXEVALG (SEQ ID NO: 20),
GX₂SX₄GX₃PX_aLXGX₂V (SEQ ID NO: 21), and
E(Y/F)X₂YX₃QX₄YLL (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid, wherein

E(Y/F)X₂YX₃QX₄YLL (SEQ ID NO: 22)

is closest to the C terminus.

47. (currently amended) An isolated poly(ADP-ribose) polymerase (PARP) homolog comprising human PARP2 (SEQ ID NO: 2) or a functional equivalent thereof which is at least 95% 85% homologous to human PARP2 (SEQ ID NO: 2) thereto, exhibits poly(ADP-ribose)-synthesizing activity, and has an amino acid sequence which

- a) has a functional NAD⁺ binding domain comprising the sequence ~~motif~~
PX_n(S/T)GX₃GKGIYFA (SEQ ID NO:11)
in which n is an integral value from 1 to 5, and the X radicals are, independently of one another, any amino acid;
- and
- b) lacks a zinc finger sequence.

48. (currently amended) The PARP homolog as claimed in claim 47, wherein said PARP lacks a zinc finger sequence ~~motif~~ of the formula
CX₂CX_mHX₂C (SEQ ID NO:30)
in which m is an integral value of 28 or 30, and
the X radicals are, independently of one another, any amino acid.

49. (currently amended) The PARP homolog as claimed in claim 47, wherein the functional

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NAD⁺ binding domain comprises the following sequence **motif**:

(S/T)XGLR(I/V)XPX_n(S/T)GX₃GKGIYFA (SEQ ID NO:12)

in which n is an integral value from 1 to 5, and

the X radicals are, independently of one another, any amino acid.

50. (currently amended) The PARP homolog as claimed in claim 47₂ wherein the functional NAD⁺ binding domain comprises the following sequence **motif**:

LLWHG(S/T)X₇IL(S/T)XGLR(I/V)XPX_n(S/T)GX₃GKGIYFAX₃SKSAXY (SEQ ID NO:13)

in which n is an integral value from 1 to 5, and

the X radicals are, independently of one another, any amino acid.

51. (currently amended) The PARP homolog as claimed in claim 47₂ further comprising a leucine zipper-like sequence:

(L/V)X₆LX₆LX₆L (SEQ ID NO: 14)

wherein X radicals are, independently of one another, any amino acid.

52. (currently amended) The PARP homolog as claimed in claim 51₂ further comprising at least one of the following [[part-]] sequences **motifs**:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15),

AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),

QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),

FYTXIPHFGX₃PP (SEQ ID NO:18), and

KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid.

53. (currently amended) The PARP homolog as claimed in claim 51₂ further comprising:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15),

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AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),
QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),
FYTXIPHXFGX₃PP (SEQ ID NO:18), and
KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid.

54. (currently amended) The PARP homolog as claimed in claim 51, further comprising:

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15),
AX₃FXKX₄KTXNXWX₅FX₃PXK (SEQ ID NO:16),
QXL(I/L)X₂IX₉MX₁₀PLGKLX₃QIX₆L (SEQ ID NO:17),
FYTXIPHXFGX₃PP (SEQ ID NO:18), and
KX₃LX₂LXDIEXAX₂L (SEQ ID NO:19)

in which the X radicals are, independently of one another, any amino acid, wherein

LX₉NX₂YX₂QLLX(D/E)X_{10/11}WGRVG (SEQ ID NO: 15)

is closest to the N terminus.

55. (currently amended) The PARP homolog as claimed in claim 47, further comprising at least one of the following:

GX₃LXEVALG (SEQ ID NO: 20),
GX₂SX₄GX₃PX_aLXGX₂V (SEQ ID NO: 21), and
E(Y/F)X₂YX₃QX₄YLL (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid.

56. (currently amended) The PARP homolog as claimed in claim 47, further comprising

GX₃LXEVALG (SEQ ID NO: 20),
GX₂SX₄GX₃PX_aLXGX₂V (SEQ ID NO: 21), and
E(Y/F)X₂YX₃QX₄YLL (SEQ ID NO: 22)

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in which a is 7 to 9 and

X is any amino acid.

57. (currently amended) The PARP homolog as claimed in claim 47, further comprising

$\text{GX}_3\text{LXEVALG}$ (SEQ ID NO: 20),

$\text{GX}_2\text{SX}_4\text{GX}_3\text{PX}_a\text{LXGX}_2\text{V}$ (SEQ ID NO: 21), and

$\text{E(Y/F)X}_2\text{YX}_3\text{QX}_4\text{YLL}$ (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid, wherein

$\text{E(Y/F)X}_2\text{YX}_3\text{QX}_4\text{YLL}$ (SEQ ID NO: 22)

is closest to the C terminus.

58. (currently amended) The PARP homolog as claimed in claim 51, further comprising at least one of the following:

GX_3LXVALG (SEQ ID NO: 20),

$\text{GX}_2\text{SX}_4\text{GX}_3\text{PX}_a\text{LXGX}_2\text{V}$ (SEQ ID NO: 21), and

$\text{E(Y/F)X}_2\text{YX}_3\text{QX}_4\text{YLL}$ (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid.

59. (currently amended) The PARP homolog as claimed in claim 51, further comprising

$\text{GX}_3\text{LXEVALG}$ (SEQ ID NO: 20),

$\text{GX}_2\text{SX}_4\text{GX}_3\text{PX}_a\text{LXGX}_2\text{V}$ (SEQ ID NO: 21), and

$\text{E(Y/F)X}_2\text{YX}_3\text{QX}_4\text{YLL}$ (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid.

60. (currently amended) The PARP homolog as claimed in claim 51, further comprising

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$\text{GX}_3\text{LXEVALG}$ (SEQ ID NO: 20),
 $\text{GX}_2\text{SX}_4\text{GX}_3\text{PX}_a\text{LXGX}_2\text{V}$ (SEQ ID NO: 21), and
 $\text{E(Y/F)X}_2\text{YX}_3\text{QX}_4\text{YLL}$ (SEQ ID NO: 22)

in which a is 7 to 9 and

X is any amino acid, wherein

$\text{E(Y/F)X}_2\text{YX}_3\text{QX}_4\text{YLL}$ (SEQ ID NO: 22

is closest to the C terminus.